

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ



**Medical Biotechnology group
School of Paramedical Sciences
Qazvin University of Medical Sciences**

Introduction to microRNAs

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Under supervision of : Dr. Ahmadpour**



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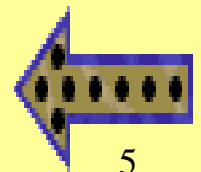
MicroRNA

- small, non-protein-coding RNA molecules~ 22 nucleotides⁽¹⁾
- **regulate** gene expression by **inhibiting** translation and/or by inducing **degradation** of target mRNAs⁽²⁾
- **Repress activity** of complementary mRNAs⁽¹⁾
- 1 miRNA = hundreds of mRNAs ⁽¹⁾



MicroRNA

- The human genome may encode over 1000 miRNAs (1)
- Regulate 60% of mammalian gene products (1)
- They have been identified to play a key role in **organ development**, **cellular differentiation**, **proliferation** and **function** (1)
- Have been described in **invertebrates** and **vertebrates**: worms, fungi, plants, and mammals (1)





History of miRs

- ❑ In **1987**, Fergunsson *et al*, found that **lin-4** could negatively regulate **lin-14**.⁽³⁾
- ❑ In **1989**, Ambros focus his research on **lin-4** gene and Ruvkun on the study **lin-14** gene.⁽⁴⁾
- ❑ In **1992** Ambros and Ruvkun : **lin-4** transcripts were complementary to a repeated sequence in the **3'UTR** of **lin-14** gene. ⁽⁵⁾



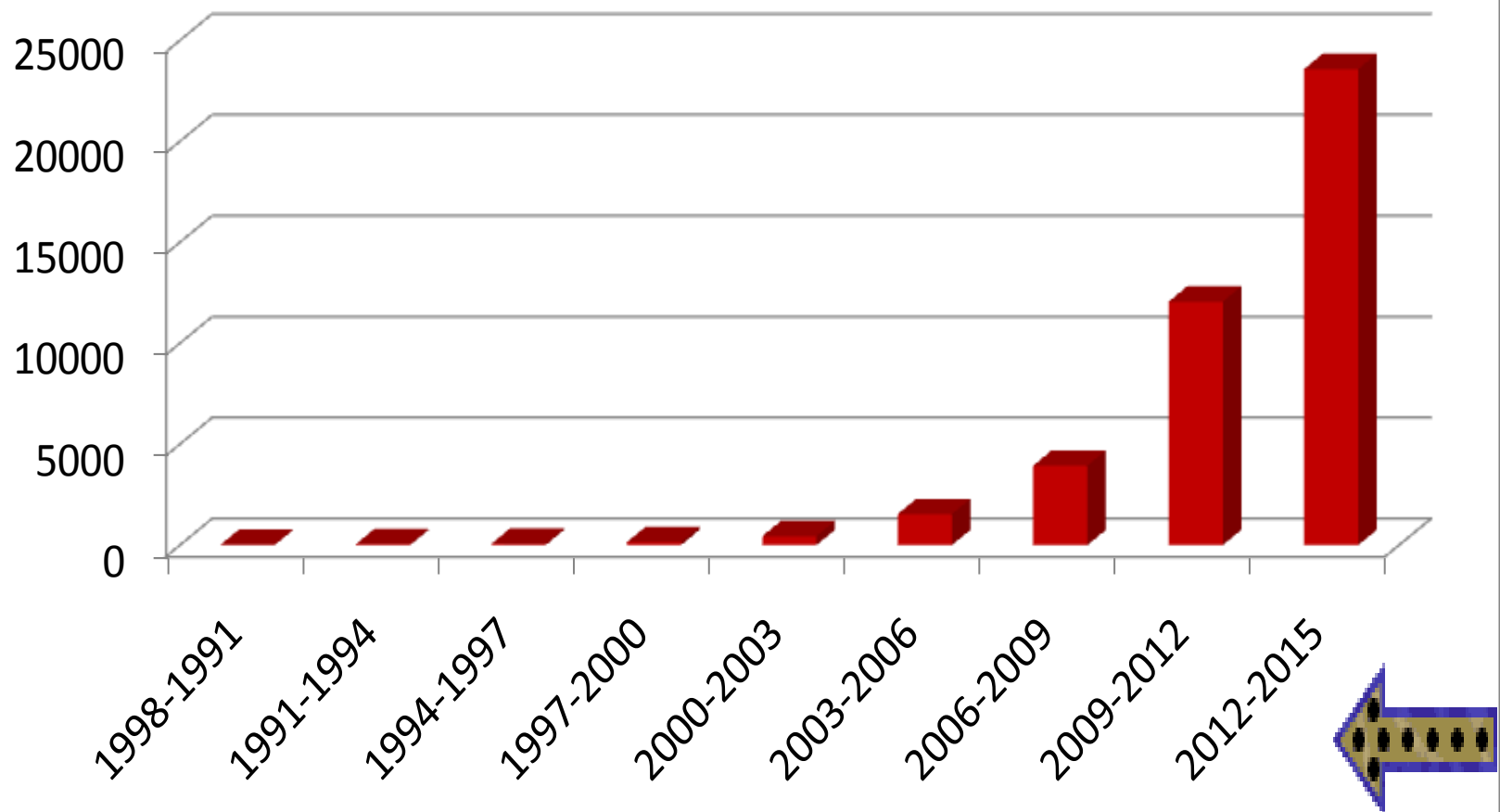
History of miRs

1993	2000	2001	2002	2003	2004
Ambros laboratory discovers the first microRNA lin-4 in <i>C. elegans</i> ¹	Ruvkun laboratory discovers let-7 microRNA in <i>C. elegans</i> ¹¹	<ul style="list-style-type: none">• Zamore laboratory shows that Dicer makes microRNA¹²• Bartel, Tuschl and Ambros laboratories discover large classes of small RNAs with regulatory roles and name them microRNAs¹³⁻¹⁵• Bartel lab discovers microRNA in plants^{16,17}	Croce laboratory shows that miRNAs are implicated in cancer because miR-15 and miR-16 are deleted or downregulated in most chronic lymphocytic leukemias ¹⁸	<ul style="list-style-type: none">• miRNA maturation shown to begin in the nucleus¹⁹• Stoffel laboratory silences miRNAs <i>in vivo</i> using chemically engineered oligonucleotides, termed 'antagomirs'²⁰	Animal viruses shown to use microRNA ²¹

Figure 1 Milestones in miRNA research.

Mack GS. Nature biotechnology. 2007;25(6):631-8.

Published Articles of MicroRNA



Biogenesis of miRNA

miRNA gene

1

pri-miRNA

2

pre-miRNA

3

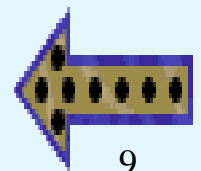
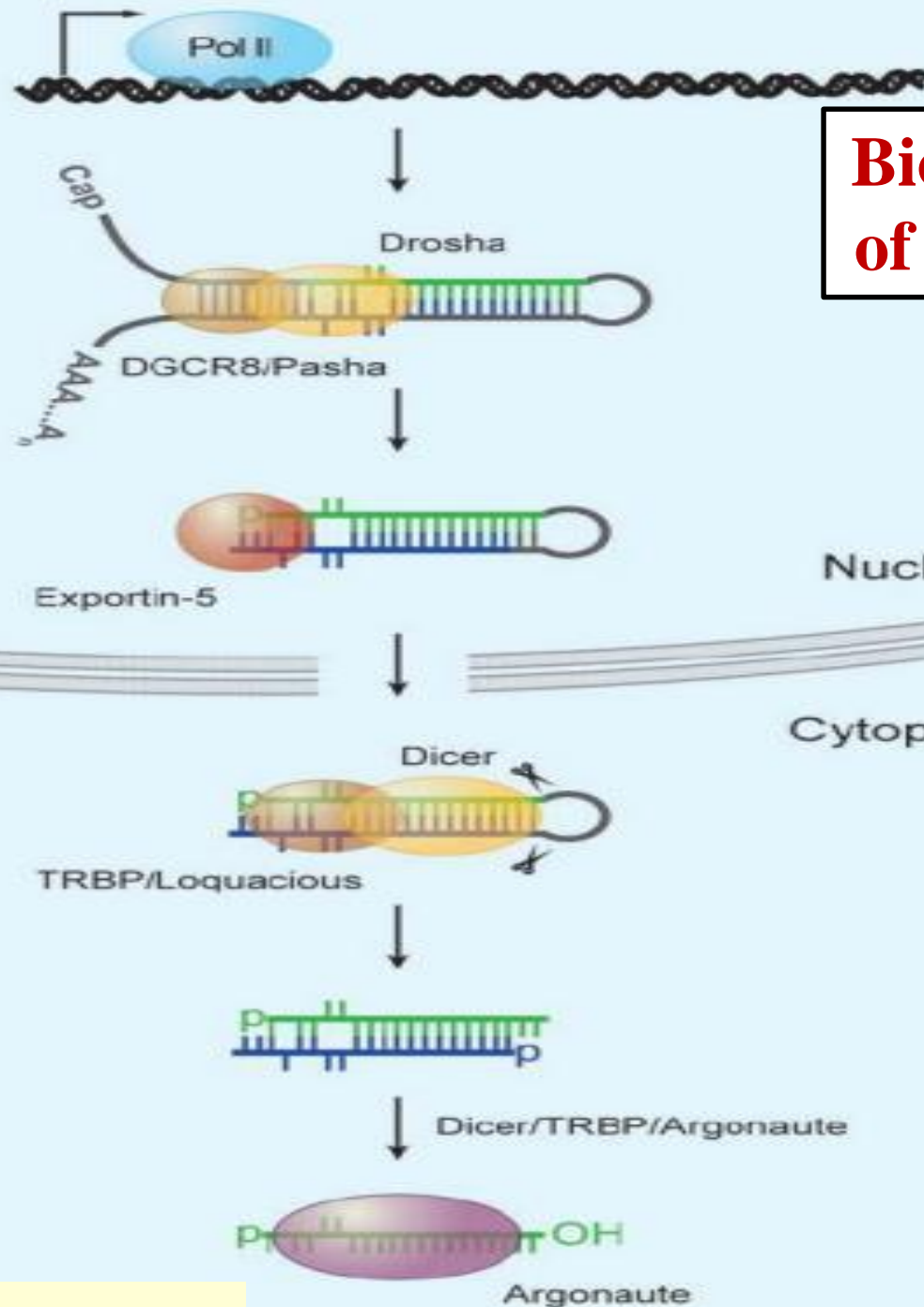
pre-miRNA

4

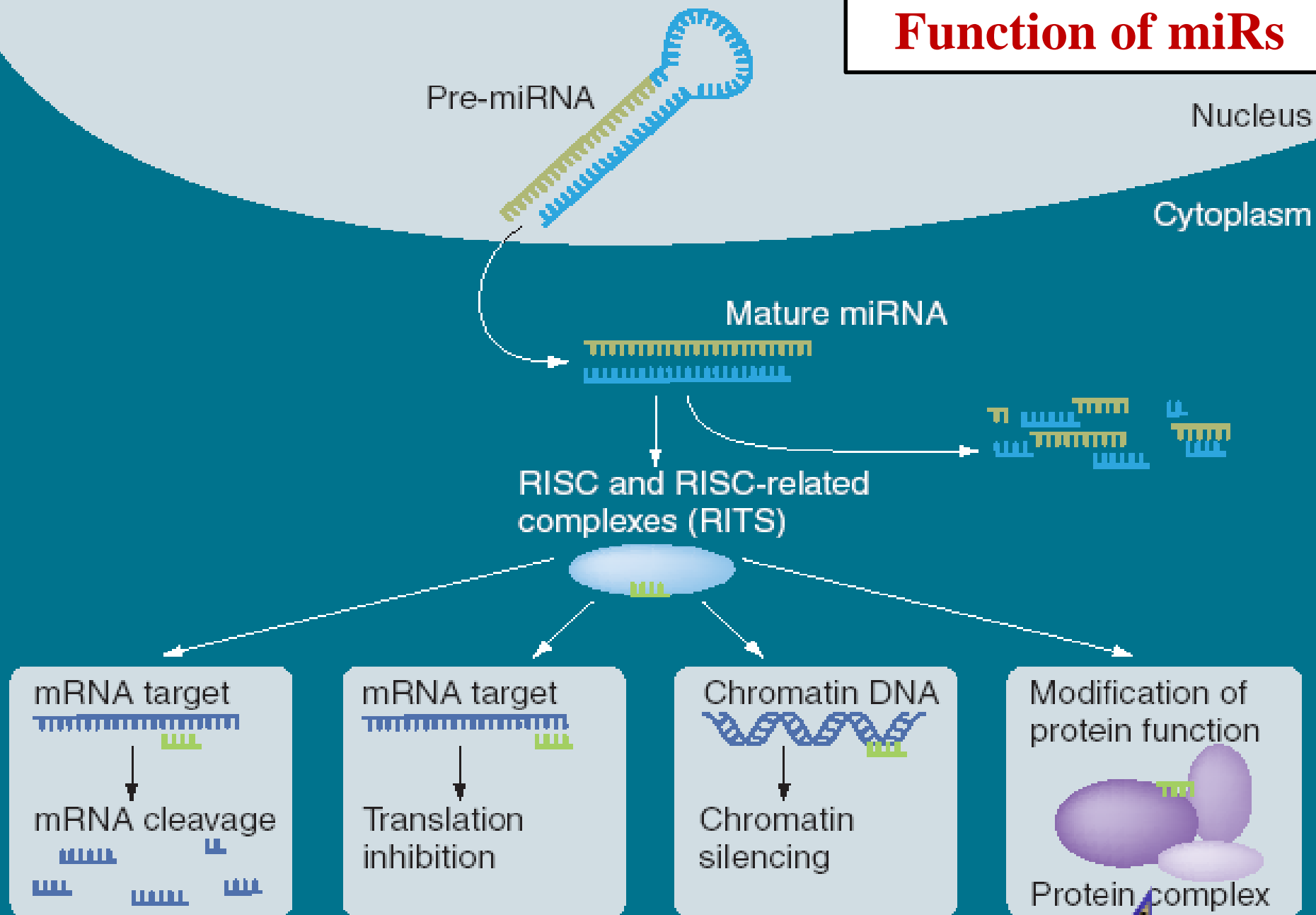
miRNA:miRNA* duplex

5

Mature miRNA
bound to Argonaute



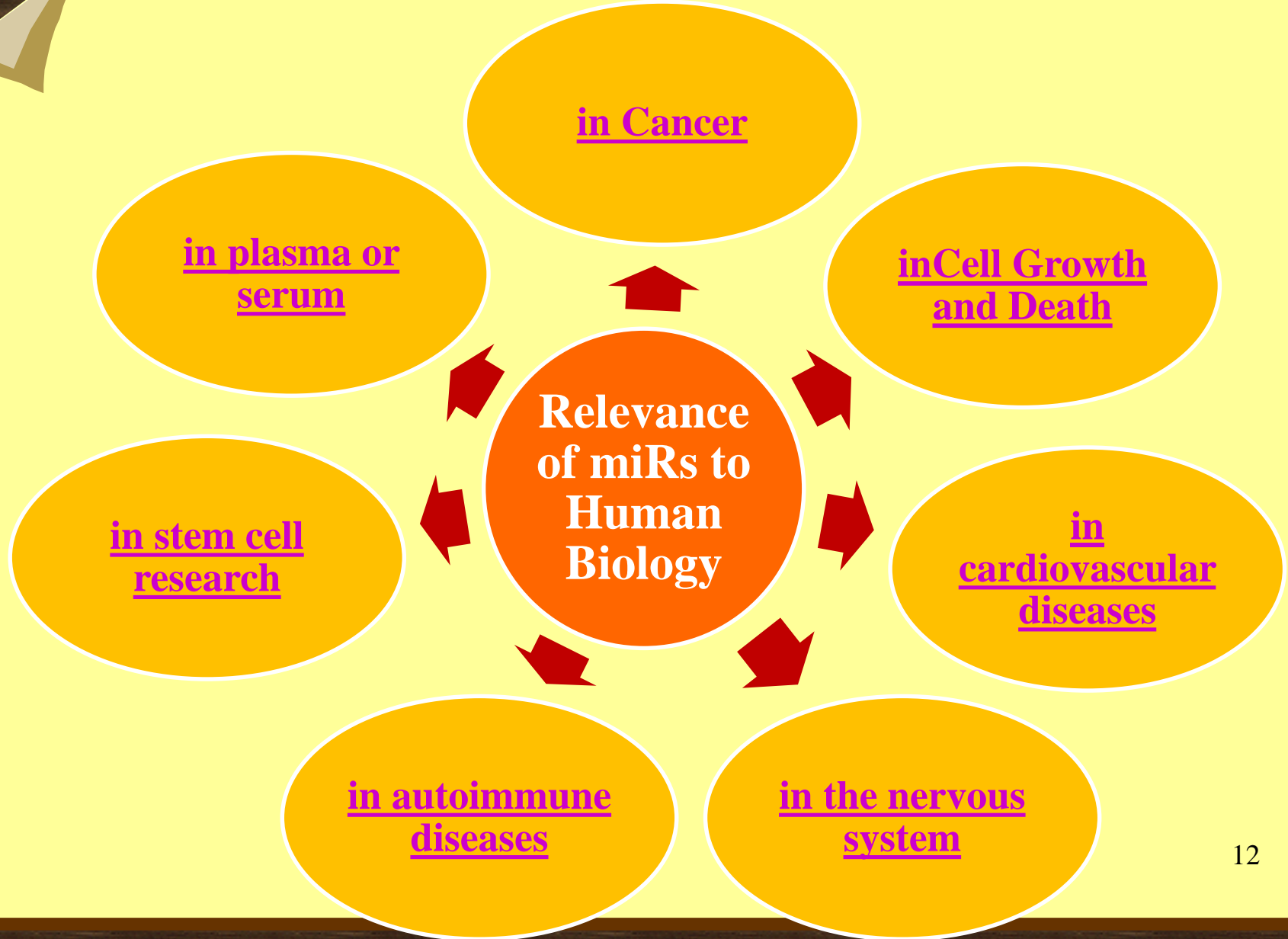
Function of miRs





Named miRs

- ✓ hsa-miR-100 → Homo sapiens
- ✓ dme-miR-100 → Drosophila melanogaster
- ✓ miR-15-5p
- ✓ miR-15-3p*
- ✓ miR-181a
- ✓ miR-181b₍₆₎

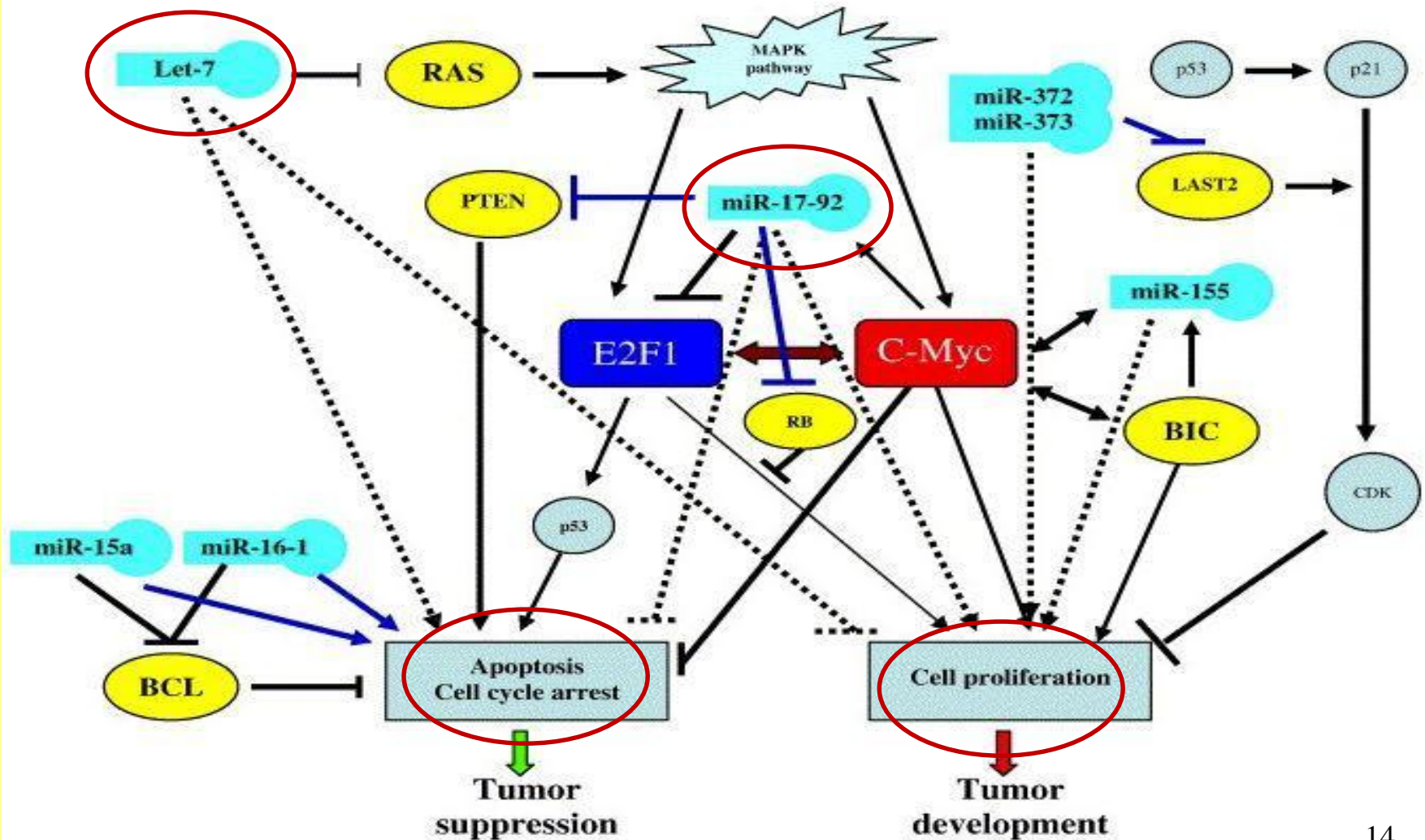




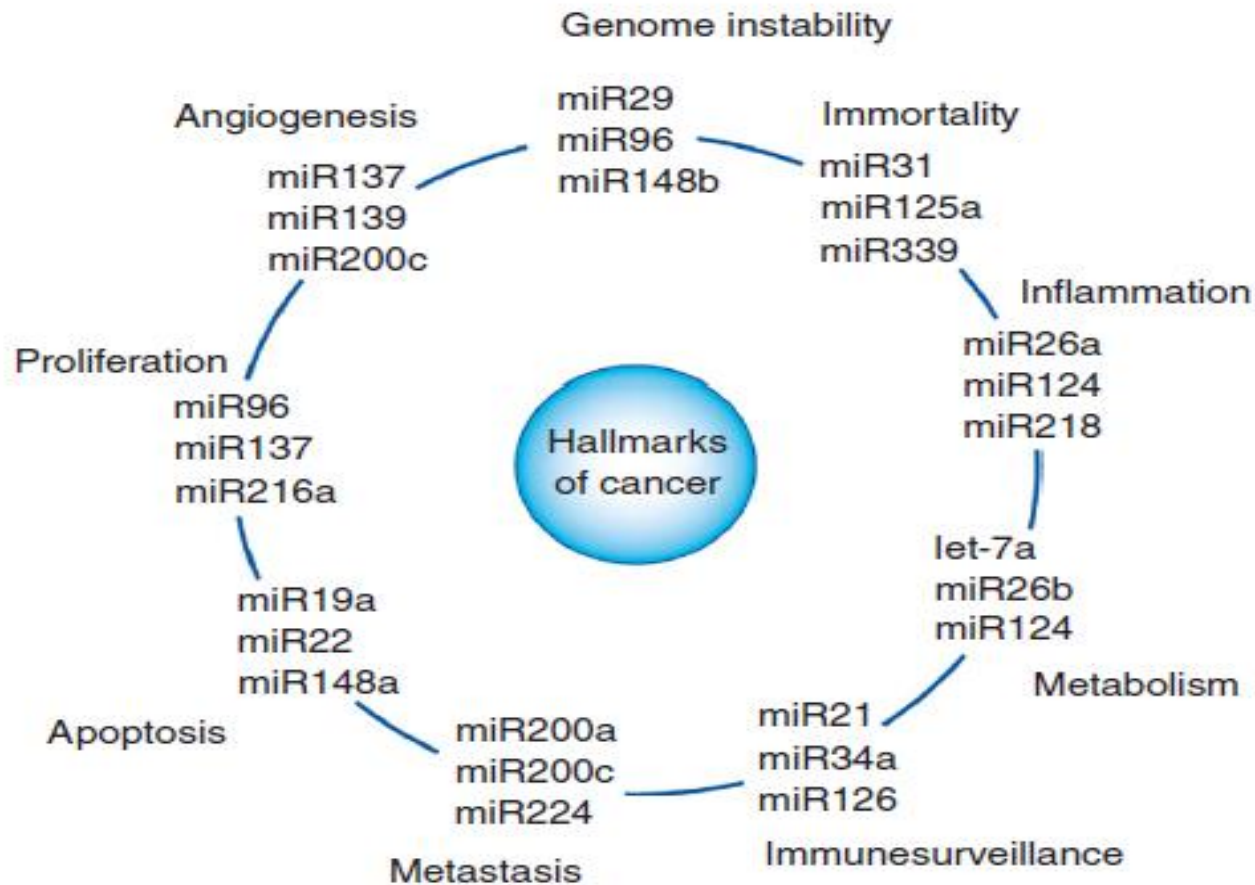
microRNAs in Cancer

- **In 2002**, the first report ,**miR-15** , **miR-16** were found to be located at chromosome **13q14**, a region frequently **deleted in CLL** (7)
- **In 2005**, MicroRNAs can act as **oncogenes** or tumor **suppressors**(8)
- **In 2006**, Saito et al. controlled by : **DNA methylation** , **histone modifications**(9)

miRNAs as Oncogenes and Tumor Suppressors



microRNAs in Cancer



Pichler M, Calin G. British journal of cancer. 2015;113(4):569-73.



microRNAs in Cancer

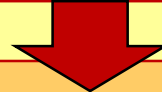
Table 1. Examples of important studies that implicated microRNAs in cancer diagnosis, prognosis and prediction of response to cancer drugs

Purpose	MicroRNA(s)	Type of cancer	Sample	Authors
Early detection	13 microRNA panel ('miR-Test')	Lung cancer	Serum	Montani <i>et al</i> , 2015
Diagnosis	24 microRNA panel	Kidney cancer	Tissue	Spector <i>et al</i> , 2013
Diagnosis	3 microRNA panel (miR21, miR31 and miR210)	Lung cancer	Sputum	Xing <i>et al</i> , 2015
Diagnosis	3 microRNA panel (miR200c, -93-3p, -192)	Mesothelioma	Tissue	Benjamin <i>et al</i> , 2010
Diagnosis	8 microRNA panel (miR106a, -125a-5p, -129-3p, -205, -21, -29b, -375, -7)	Lung cancer	Tissue	Gilad <i>et al</i> , 2012
Diagnosis	64 microRNA panel	Cancer of unknown primary	Tissue	Meiri <i>et al</i> , 2012
Diagnosis	48 microRNA panel	Cancer of unknown primary	Tissue	Rosenfeld <i>et al</i> , 2008
Prognosis	miR224	Colorectal cancer	Tissue	Ling <i>et al</i> , 2015
Prognosis	miR200a	Colorectal cancer	Tissue	Pichler <i>et al</i> , 2014
Prognosis	let-7i, miR-10b, miR-885-5p	Colorectal cancer	Tissue	Hur <i>et al</i> , 2015
Prognosis	miR155	Lymphoma	Cells	Iqbal <i>et al</i> , 2015
Prediction	miR212	Head and neck cancer	Tissue	Hatakeyama <i>et al</i> , 2010
Prediction	let7-binding site in KRAS gene	Colorectal cancer	Tissue or blood	Zhang <i>et al</i> , 2011

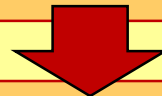


microRNAs in Cancer

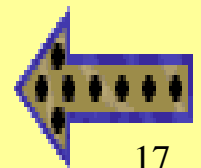
in 2015



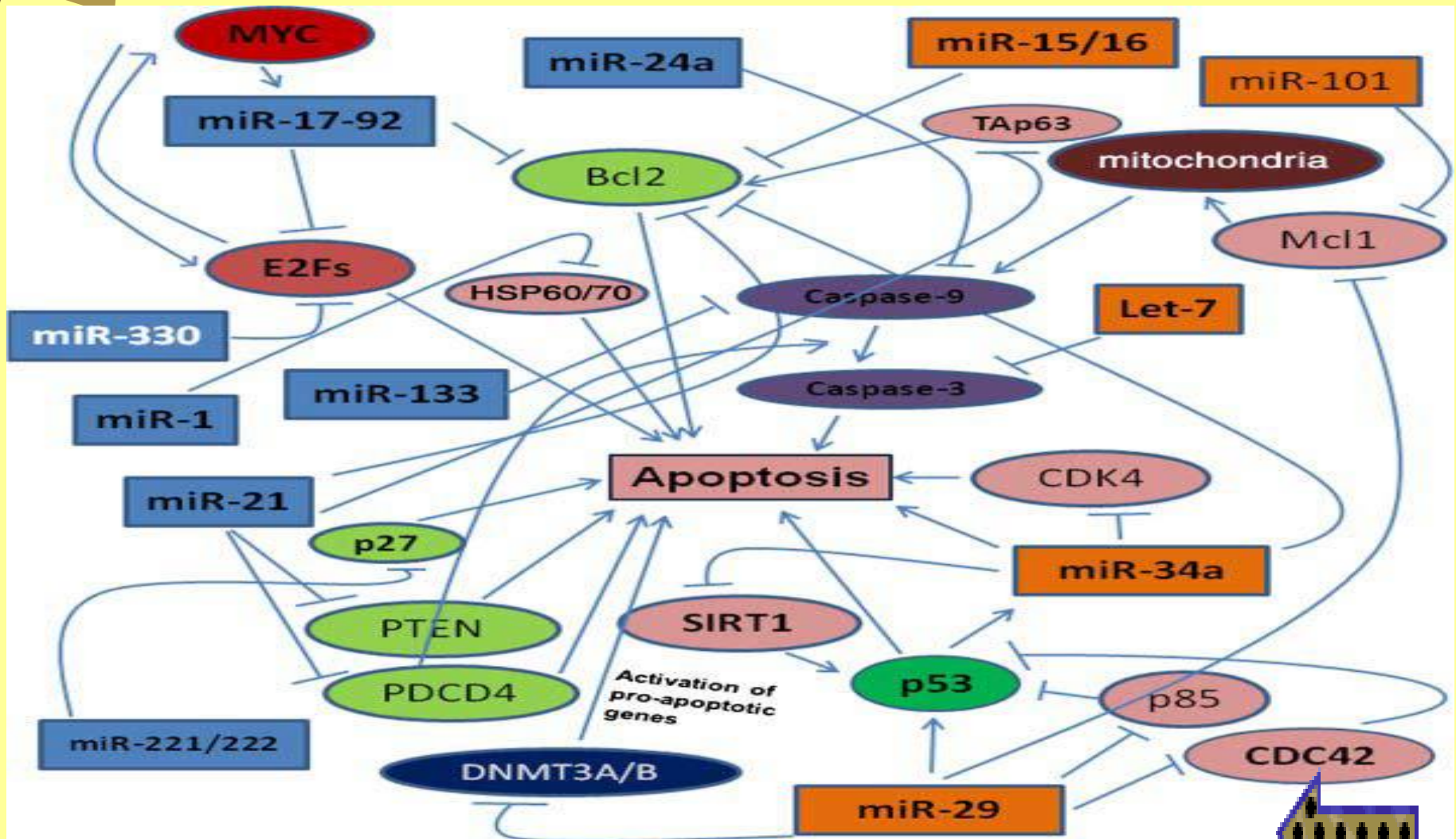
Ferracin M et al



miR-181a-5p as a diagnostic
breast cancer biomarker⁽¹⁰⁾.



miRNAs Regulate Cell Growth and Death

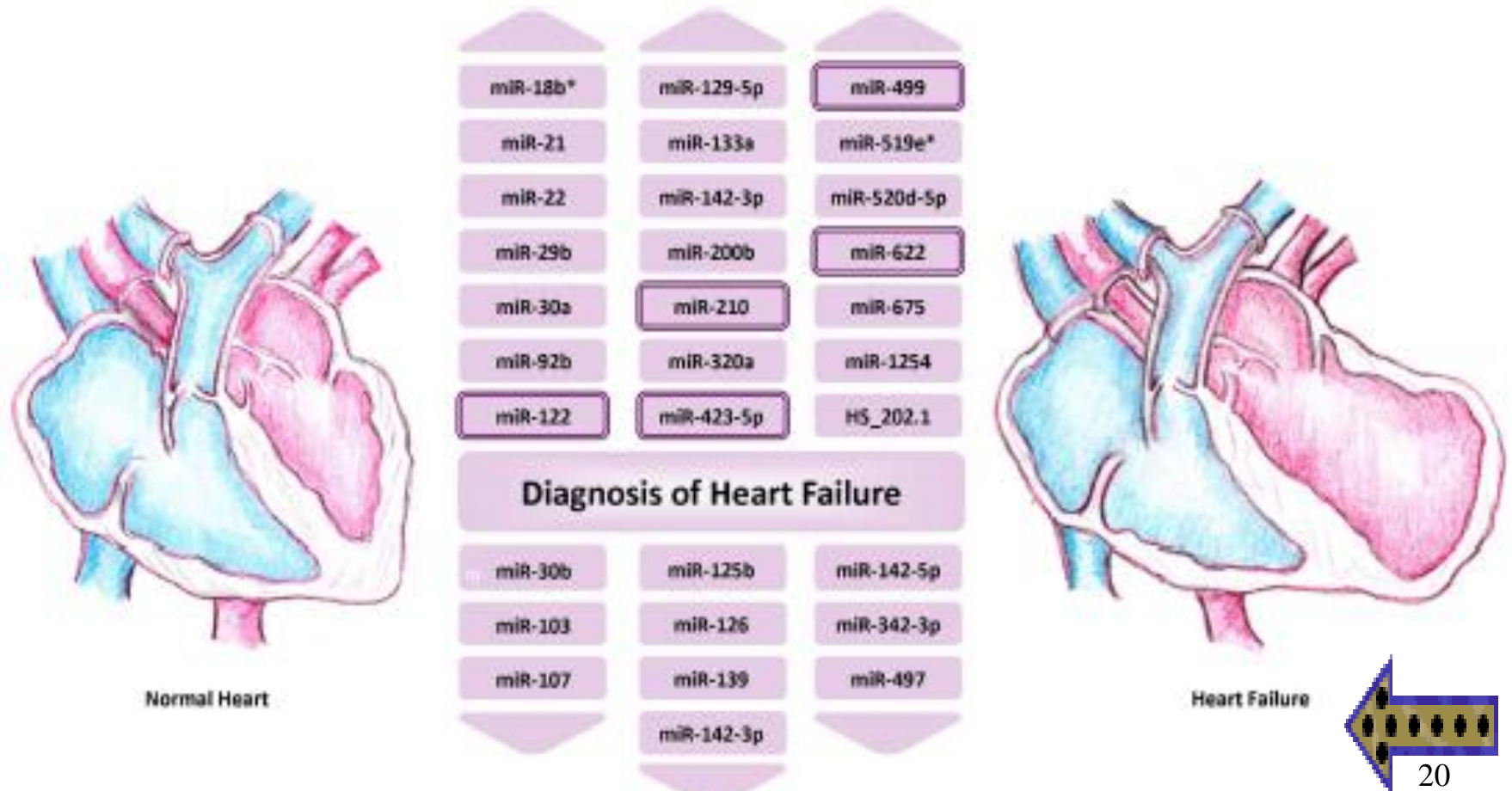




microRNAs in cardiovascular diseases

- In 2005, Zhao et al. miR-1-1, miR-1-2 were specifically expressed in cardiac and skeletal muscle precursor cells⁽¹¹⁾.
- In 2006, Rooij et al. microarray analysis, more than 12 miRNAs were cardiac hypertrophy and heart failure ⁽¹²⁾.
- In 2010, Edwards et al. several studies demonstrated miRNA aberrant patterns in cardiac hypertrophy and analyzed their roles⁽¹³⁾.

microRNAs in cardiovascular diseases

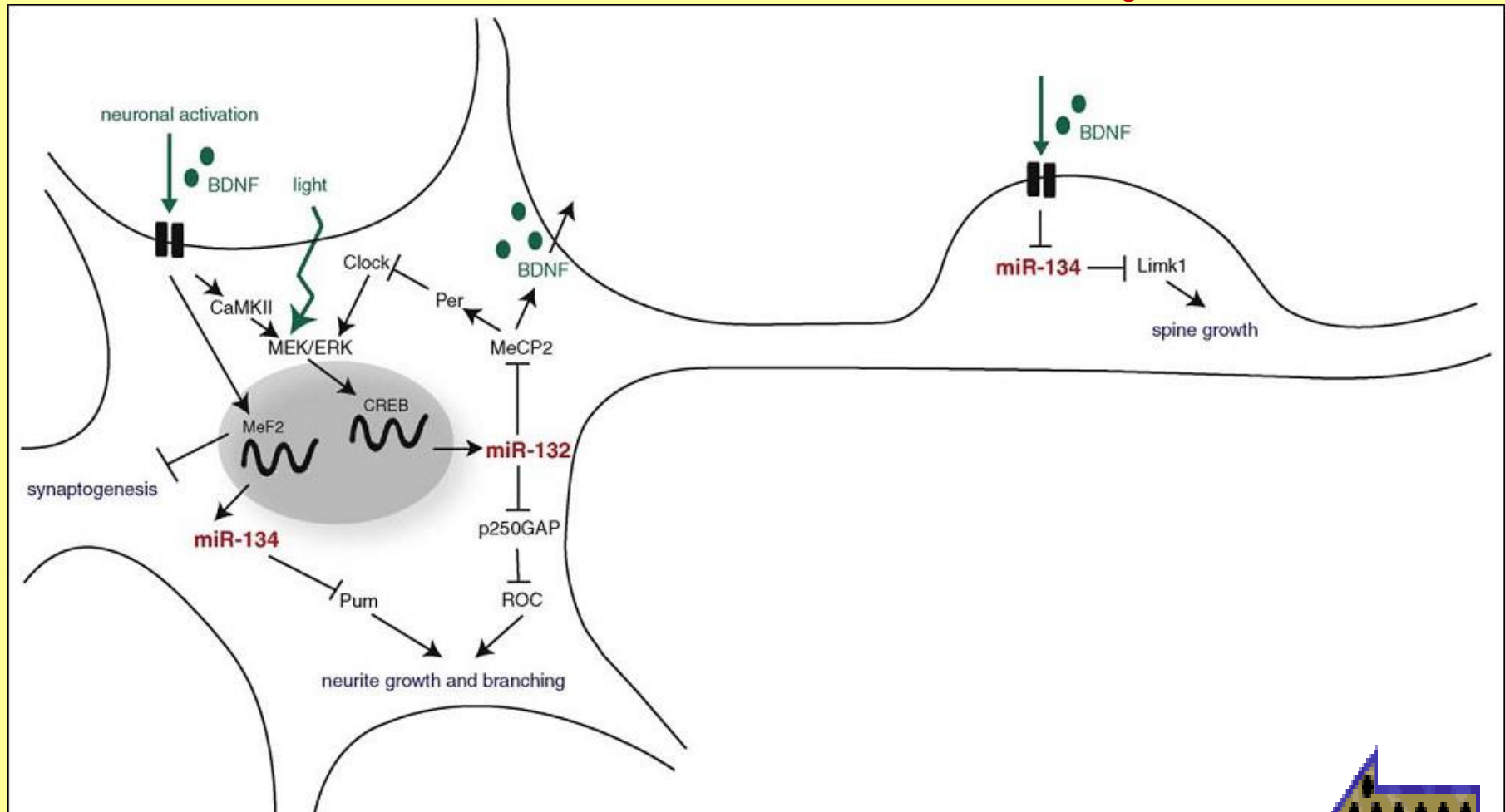




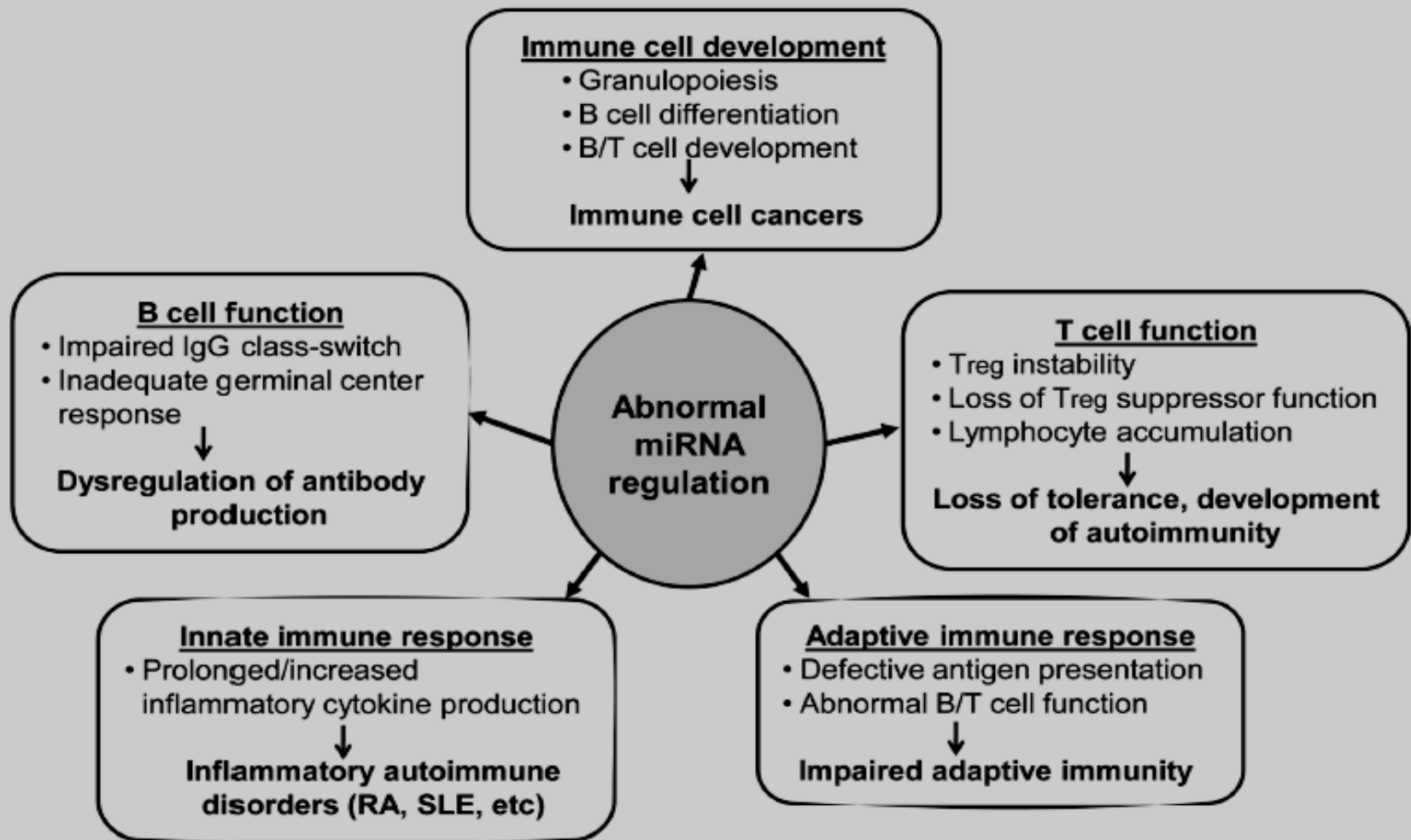
microRNAs in the nervous system

- key regulators of CNS development
 - involving brain plasticity
 - neuronal cell maturation
 - neuronal cell differentiation
 - synaptogenesis
- Dysregulation ➡ Alzheimer, Parkinson, Huntington (14)

microRNAs in the nervous system



microRNAs in autoimmune diseases

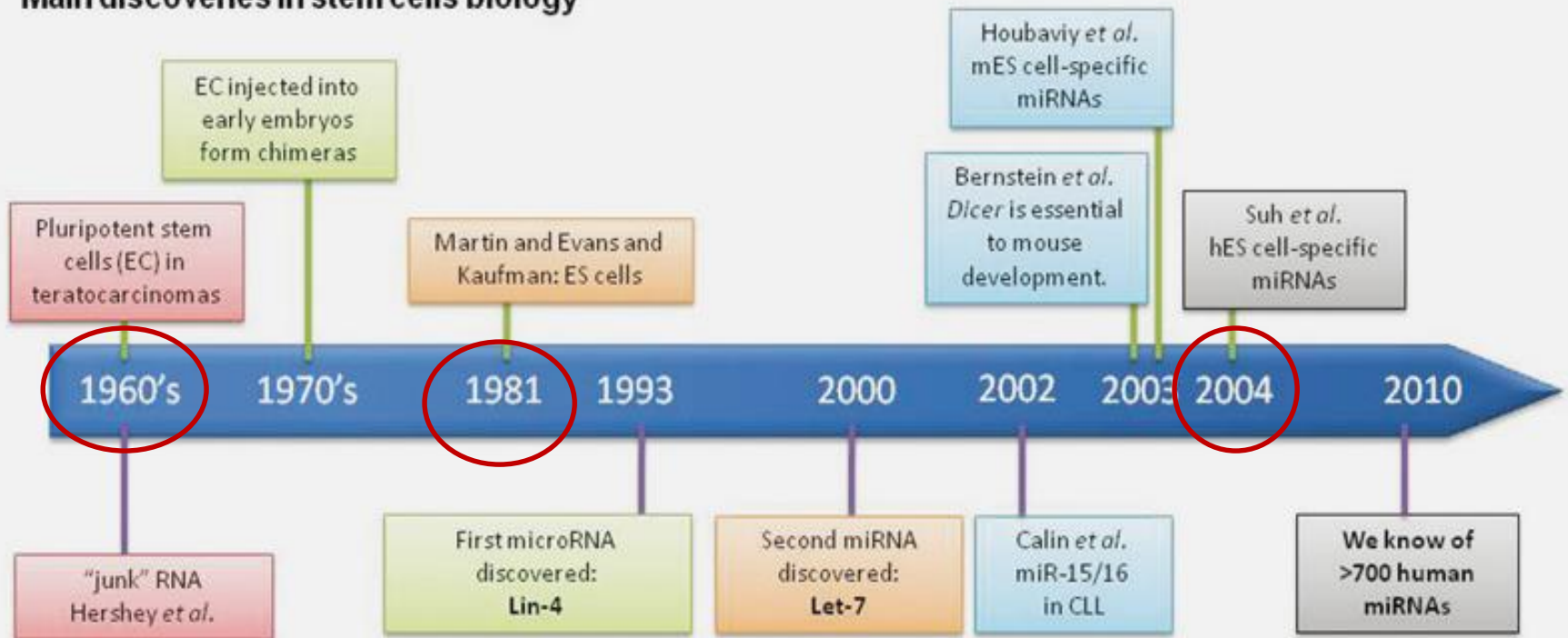


microRNAs in autoimmune diseases

Diseases	Serum miRNAs	Expression levels
SLE	MiR-146a	↓
	MiR-155	↓
	MiR-200a	↓
	MiR-342-3p, MiR-223, and MiR-20a	↓
	MiR-142-3p	↓
	MiR-126	↑
PS	MiR-369-3p	↑
SSc	MiR-29a	↓
	MiR-196a	↓
	MiR-92a	↑
	MiR-150	↓
DM	MiR-7,223	↓

microRNAs in stem cell research

Main discoveries in stem cells biology



Main discoveries in microRNAs biology



microRNAs in stem cell research

In 2015 Zhou C et al

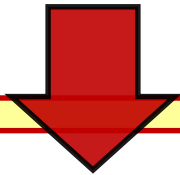
overexpression of pluripotency stem cell-specific **miRNA-302/367** cluster, **miRNA-9/9** and **miRNA-124** induced fibroblasts conversion into neurons.

important implications : studies of neuron development and neurological disease therapy.(15)

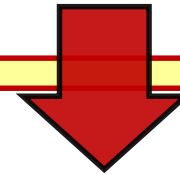


microRNAs in stem cell research

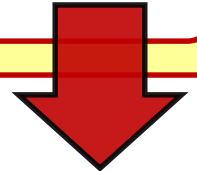
In 2015 Möbus S et al



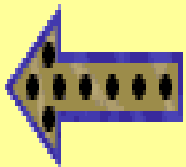
miR-199a-5p inhibition in ES-derived HLCs leads to improved hepatocyte differentiation.



Upon transplantation, HLCs were able to engraft and repopulate the liver of mice.



miRNA modulation may serve as a promising approach to generate more mature HLCs from stem cell sources for the treatment of liver diseases.(16)





microRNAs as biomarkers in plasma or serum

in 2008 first report

Chim et al

miRNAs as diagnostic tools in
biological fluids

detected placental miRNAs in the
maternal plasma⁽¹⁷⁾

in 2008 Lawrie et al

comparing serum from diffuse large
B-cell lymphoma patients and
healthy controls

miR-155, miR-210 and miR-21
levels were significantly upregulated
in patients⁽¹⁸⁾



microRNAs as biomarkers in plasma or serum

in 2010 Murata et al

analyzed miRNAs plasma levels in
rheumatoid arthritis

plasma **miR-132** was significantly
higher in healthy controls than in
rheumatoid arthritis⁽¹⁹⁾

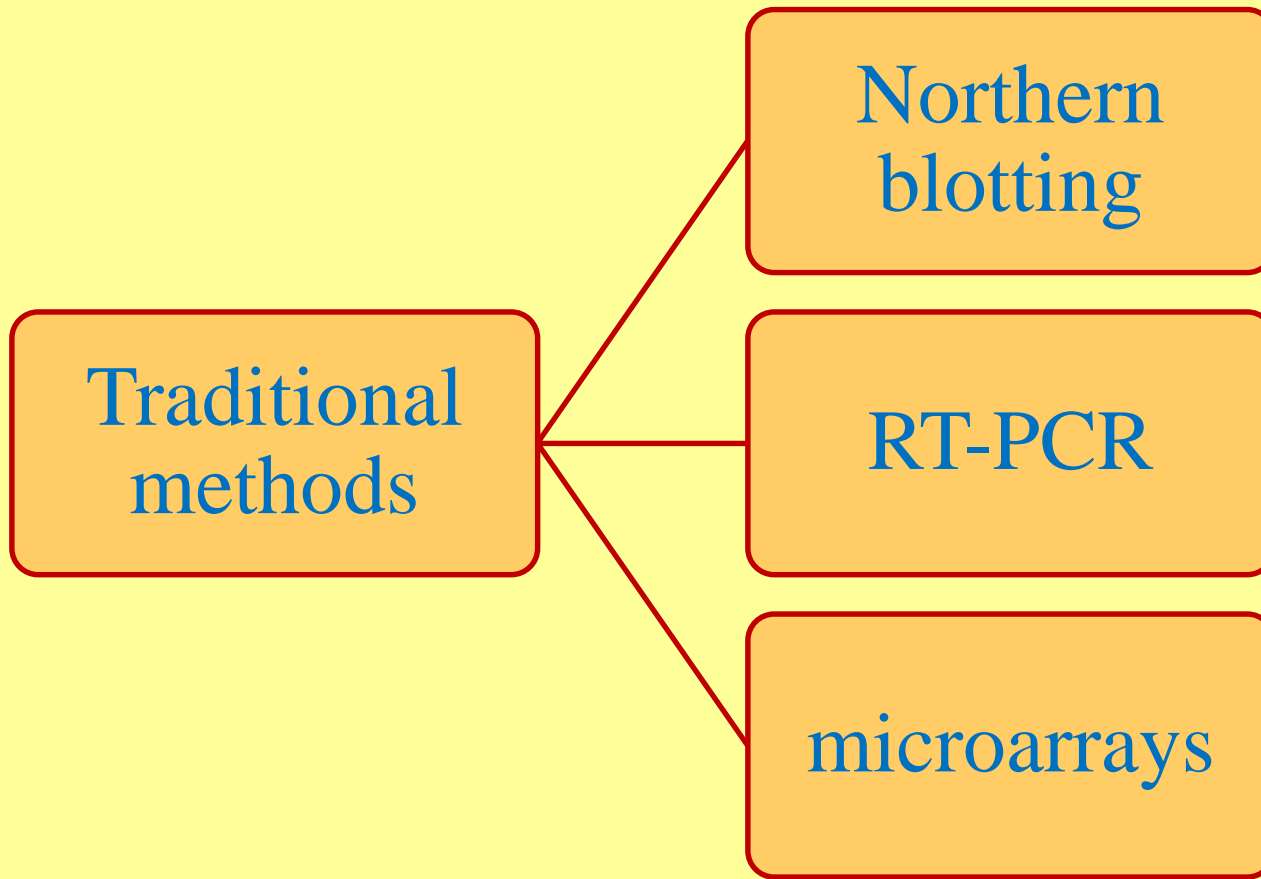
in 2015 Tsochandaridis et
al

noninvasive diagnostic and
prognostic biomarkers for pregnancy
monitoring

prevent the development of
gestational disorders and form the
basis of personalized therapeutic
strategies⁽²⁰⁾

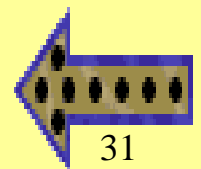
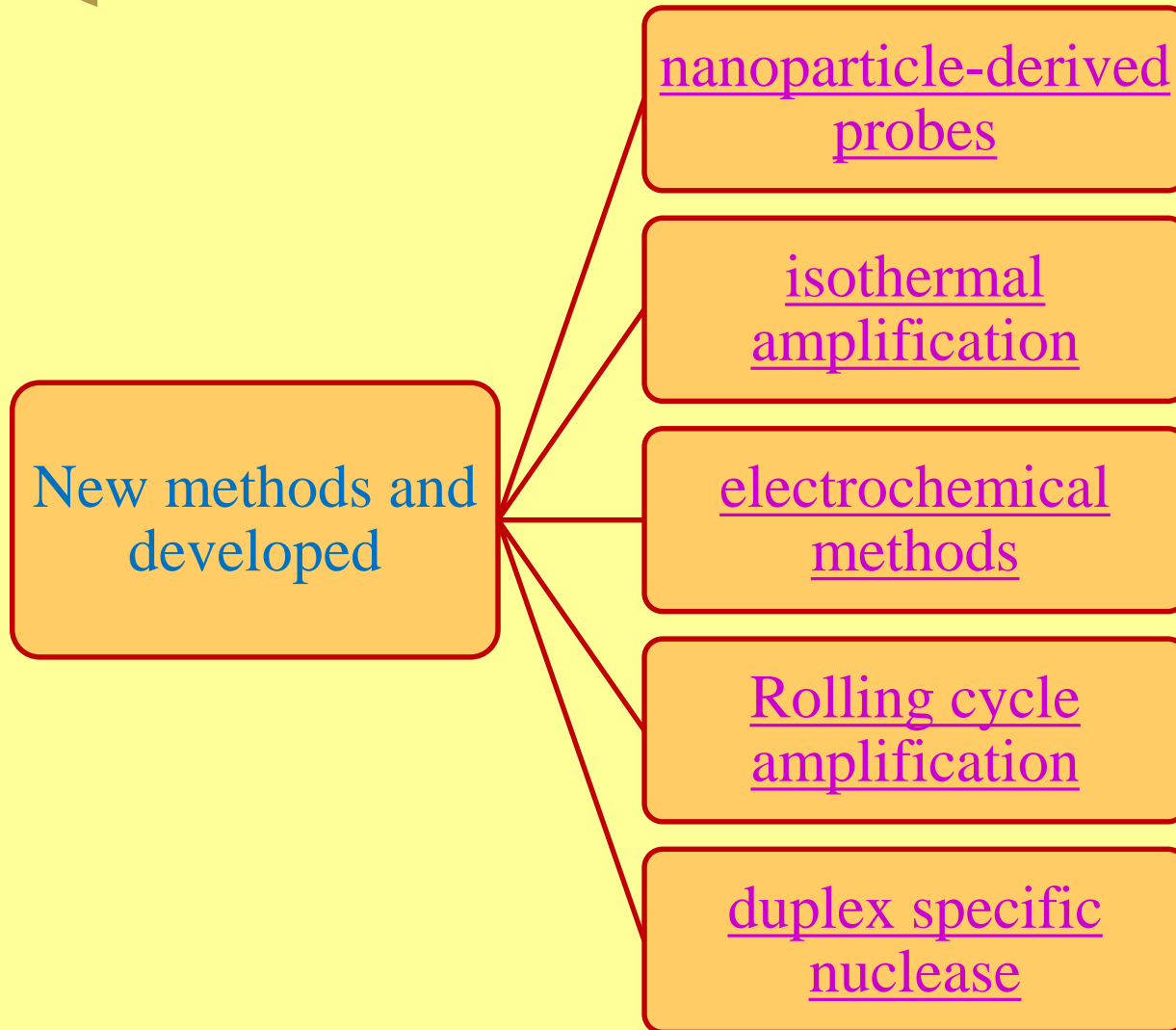


microRNA detection methods⁽²¹⁾





microRNA detection methods





nanotechnology and nanomaterials

- potent tools in bioanalysis
- detection of biological targets
- high sensitivity
- high efficiency

Nanoparticles



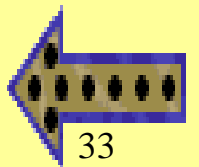
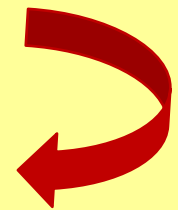


AuNPs-based methods

- ✓ Simple probe design,
- ✓ easy modification,
- ✓ biocompatibility,
- ✓ easy cellular-uptake

Disadvantages

Specialized nanotechnology platform needed





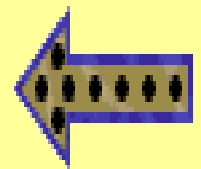
Quantum dot-based methods

- ✓ Special fluorescence properties
- ✓ also easy modification
- ✓ simple design of probe
- ✓ easy cellular-uptake

Disadvantages

Cell cytotoxicity

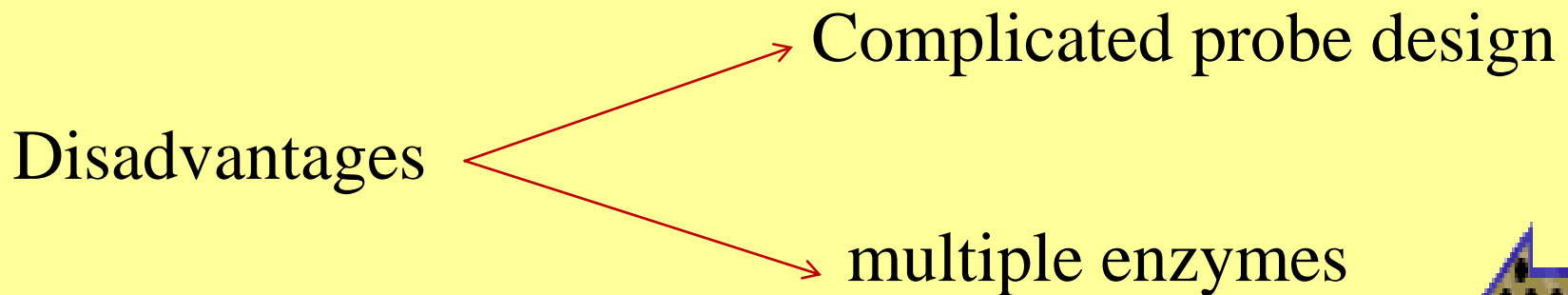
unsuitable for multiple target detection





Isothermal exponential amplification


- ✓ No PCR needed
- ✓ good efficiency of signal amplification
- ✓ good sensitivity even low at the single cell level
- ✓ real-time assay in some cases





Some cleavage-based methods

- ✓ Simplified enzymatic system
- ✓ good sensitivity and selectivity
- ✓ less side reactions

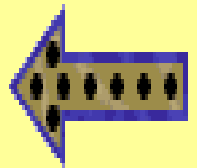
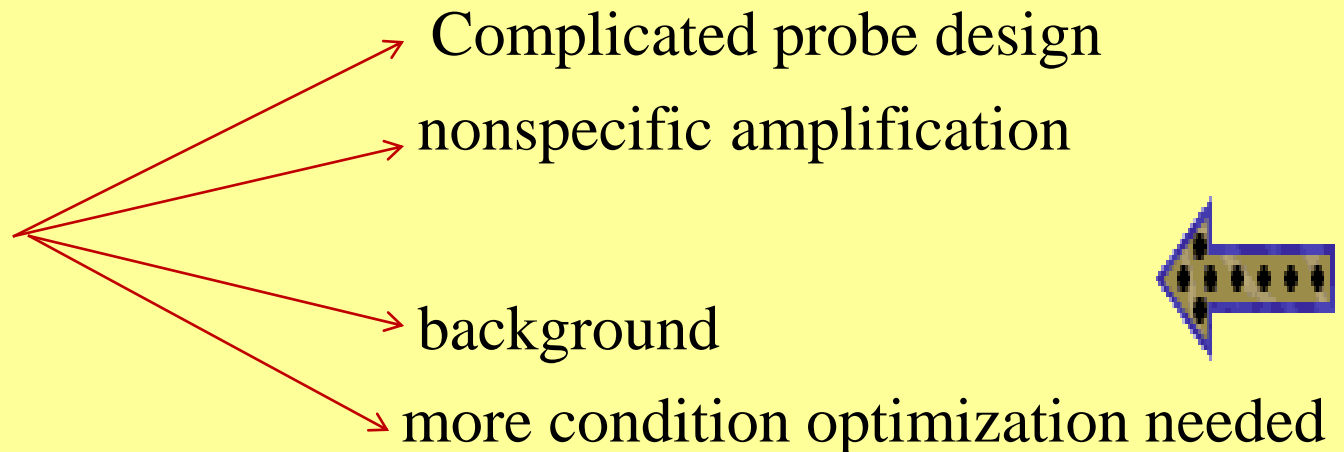
Disadvantages  Special enzymes needed



Rolling cycle amplification

- ✓ no PCR needed,
- ✓ good efficiency of signal amplification
- ✓ in situ detection achieved in some cases

Disadvantages





DSN and AuNPs-based system

- ✓ 0.2 fmol level detection
- ✓ simple design of probe

Disadvantages  Time consuming

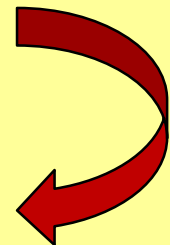


Capillary-electrophoresis-based assay

- ✓ Simple probe design
- ✓ multiple target detection

Disadvantages

Specialized equipment need





Using bioinformatics tools to identify miRs

- ➡ KEGG: [http:// www.genome.jp/kegg/](http://www.genome.jp/kegg/)
- ➡ Target Scan : <http://www.targetscan.org/>
- ➡ miRBase: <http://www.mirbase.org/>
- ➡ miRWalk: <http://www.umm.uni-heidelberg.de/apps/zmf/mirwalk>
- ➡ miRanda :<http://www.microrna.org>



Using bioinformatics tools to identify miRs

- ➡ Diana-microT: <http://diana.cslab.ece.ntua.gr/>
- ➡ TarBase: : <http://diana.cslab.ece.ntua.gr/>
- ➡ miRDB : <http://mirdb.org/>
- ➡ GEO: www.ncbi.nlm.nih.gov/gds
- ➡ miRNAMap: <http://mirnamap.mbc.nctu.edu.tw/>



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KEGG: Kyoto Encyclopedia of Genes and Genomes

KEGG is a database resource for understanding high-level functions and utilities of the biological system, such as the cell, the organism and the ecosystem, from molecular-level information, especially large-scale molecular datasets generated by genome sequencing and other high-throughput experimental technologies (See [Release notes](#) for new and updated features).

New article - [KEGG as a reference resource for gene and protein annotation](#)

● Main entry point to the KEGG web service

[KEGG2](#)

[KEGG Table of Contents](#)

[Update notes](#)

● Data-oriented entry points

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[Health information resource](#) [[Drug labels search](#)]

● Organism-specific entry points

[KEGG Organisms](#)

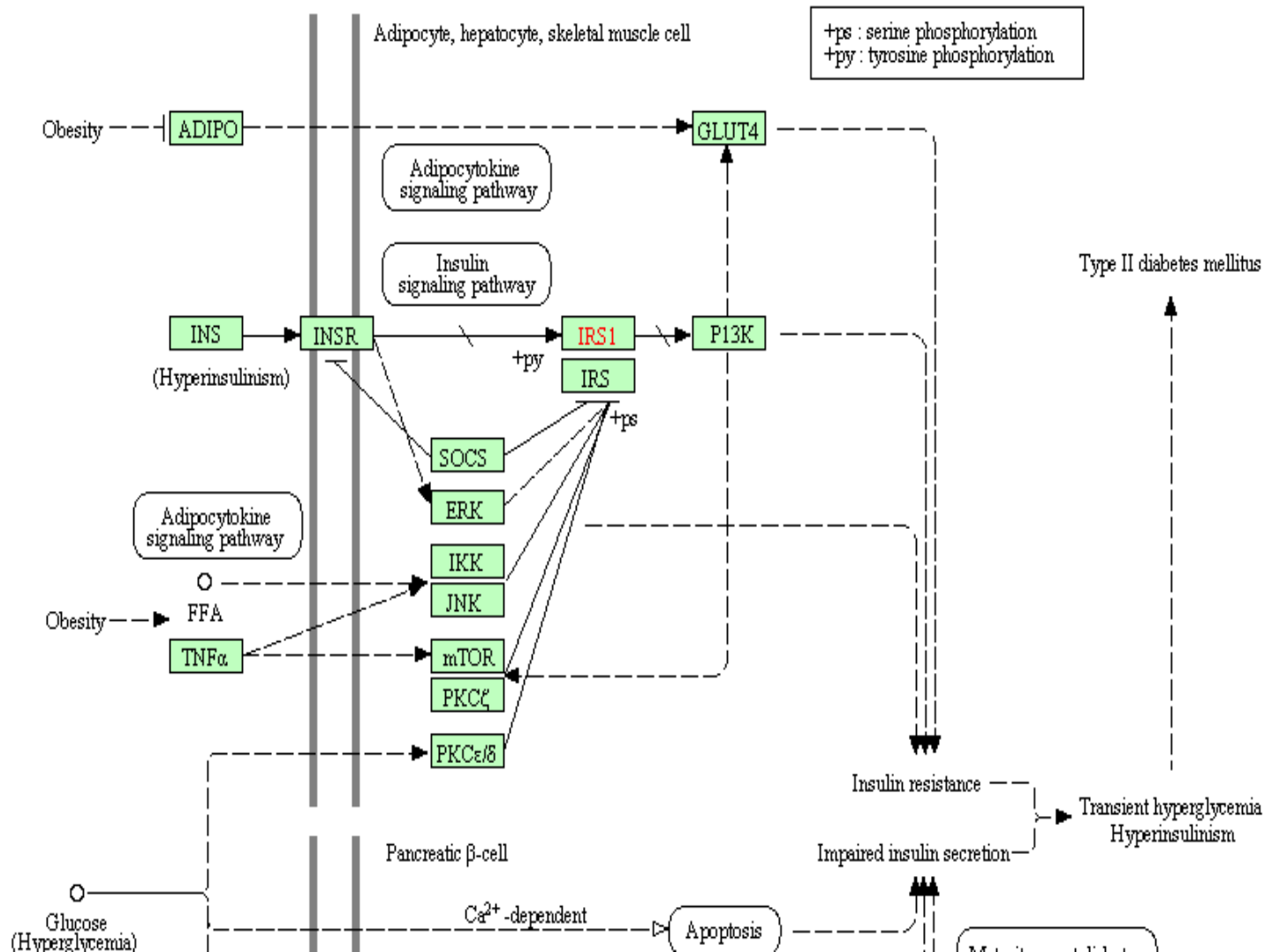
Enter org code(s)

Go

[hsa](#)

[hsa eco](#)

TYPE II DIABETES MELLITUS



Search for predicted microRNA targets in mammals

[\[Go to TargetScanMouse\]](#)

[\[Go to TargetScanWorm\]](#)

[\[Go to TargetScanFly\]](#)

[\[Go to TargetScanFish\]](#)

1. Select a species

AND

2. Enter a human gene symbol (e.g. "HMGA2")
or an Ensembl gene (ENSG00000149948) or transcript (ENST00000403681) ID

AND/OR

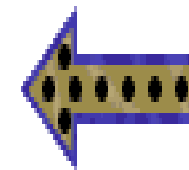
3. Do one of the following:

• Select a broadly conserved* microRNA family

• Select a conserved* microRNA family

• Select a poorly conserved microRNA family Note that many of these families are star miRNAs or small RNAs that have been misclassified as miRNAs.

• Enter a microRNA name (e.g. "miR-1-3p")



Human | miR-135-3p

100 transcripts with sites, containing a total of 160 sites.

Top 100 predicted targets, irrespective of site conservation, ranked by cumulative weighted context++ score.

The table shows at most one transcript per gene, selected for being the most prevalent, based on 3P-seq tags (or the one with the longest 3' UTR, in case of a tie). [\[Download table\]](#)

Target gene	Representative transcript	Gene name	Number of 3P-seq tags supporting UTR + 5	Link to sites in UTRs	Site counts				6mer sites	Representative miRNA	Cumulative weighted context++ score	Total context++ score	Aggregate P _{CT}
					total	8mer	7mer-m8	7mer-A1					
ZNF791	ENST00000446165.1	zinc finger protein 791	242	Sites in UTR	5	0	0	5	5	hsa-miR-135a-3p	-1.43	-1.48	N/A
C15orf53	ENST00000318792.1	chromosome 15 open reading frame 53	5	Sites in UTR	3	0	3	0	0	hsa-miR-135a-3p	-0.83	-0.83	N/A
C4orf40	ENST00000344526.5	chromosome 4 open reading frame 40	5	Sites in UTR	3	1	2	0	0	hsa-miR-135a-3p	-0.82	-0.82	N/A
ZNF629	ENST00000262525.4	zinc finger protein 629	44	Sites in UTR	4	0	0	4	3	hsa-miR-135a-3p	-0.80	-0.80	N/A
ADO	ENST00000373783.1	2-aminoethanethiol (cysteamine) dioxygenase	686	Sites in UTR	2	1	1	0	0	hsa-miR-135a-3p	-0.78	-0.78	N/A
ZNF763	ENST00000343949.5	zinc finger protein 763	5	Sites in UTR	1	0	0	1	7	hsa-miR-135a-3p	-0.75	-0.75	N/A
AC069547.1	ENST00000595931.1	HCG1745369; PRO3073; Uncharacterized protein	5	Sites in UTR	1	1	0	0	0	hsa-miR-135a-3p	-0.74	-0.74	N/A
ANAPC5	ENST00000441917.2	anaphase promoting complex subunit 5	1380	Sites in UTR	1	1	0	0	1	hsa-miR-135a-3p	-0.73	-0.74	N/A
ZNF214	ENST00000278314.4	zinc finger protein 214	7	Sites in UTR	2	0	0	2	1	hsa-miR-135a-3p	-0.70	-0.70	N/A
UPK2	ENST00000264031.2	uroplakin 2	5	Sites in UTR	1	1	0	0	0	hsa-miR-135a-3p	-0.70	-0.70	N/A
PAPPA-AS1	ENST00000445861.2	PAPPA antisense RNA 1	5	Sites in UTR	2	0	2	0	0	hsa-miR-135a-3p	-0.70	-0.70	N/A
ZNF23	ENST00000497160.1	zinc finger protein 23	5	Sites in UTR	1	0	0	1	7	hsa-miR-135a-3p	-0.69	-0.69	N/A
KRT12	ENST00000251643.4	keratin 12	5	Sites in UTR	2	0	2	0	0	hsa-miR-135a-3p	-0.69	-0.69	N/A
LIN7C	ENST00000278193.2	lin-7 homolog C (C. elegans)	1701	Sites in UTR	3	0	1	2	0	hsa-miR-135a-3p	-0.68	-0.68	N/A

Latest miRBase blog posts

[High confidence miRNA set available for miRBase 21](#)

By [sam](#) (July 3, 2014)

As mentioned previously, we briefly held off from releasing the set of "high confidence" miRNAs for miRBase 21, because of a last-gasp bug. Those data are now available, tagged with the label "high confidence" on the entry pages, and for download on the FTP site. The total number of miRNAs labelled "high confidence" has increased [...]

[miRBase 21 finally arrives](#)

By [sam](#) (June 26, 2014)

Apologies for the longer-than-usual wait. miRBase 21 is now available on the website, and all data available for download on the FTP site. As usual, the release notes describe the major changes. Of particular note this time, the Genome Reference Consortium have released a new human genome assembly, GRCh38. We have therefore remapped the human [...]

miRNA count: 28645 entries

[Release 21](#): June 2014

Search by miRNA name or keyword

Download published miRNA data

[Download page](#) | [FTP site](#)

[Tweets by @mirbase](#)

miRBase: the microRNA database

miRBase provides the following services:

- The [miRBase database](#) is a searchable database of published miRNA sequences and annotation. Each entry in the miRBase Sequence database represents a predicted hairpin portion of a miRNA transcript (termed mir in the database), with information on the location and sequence of the mature miRNA sequence (termed miR). Both hairpin and mature sequences are available for [searching](#) and [browsing](#), and entries can also be retrieved by name, keyword, references and annotation. All sequence and annotation data are also [available for download](#).
- The [miRBase Registry](#) provides miRNA gene hunters with unique names for novel miRNA genes prior to publication of results. Visit the [help pages](#) for more information about the naming service.

To receive email notification of data updates and feature changes please subscribe to the [miRBase announcements mailing list](#). Any queries about the website or naming service should be directed at mirbase@manchester.ac.uk.

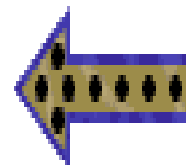
miRBase is managed by the [Griffiths-Jones lab](#) at the [Faculty of Life Sciences, University of Manchester](#) with funding from the [BBSRC](#). miRBase was previously hosted and supported by the [Wellcome Trust Sanger Institute](#).

Accession	ID	miRNA name	Previous ID	Mature name	Previous Mature ID
MI0000161	mmu-mir-135a-1	✓			
MI0000452	hsa-mir-135a-1	✓			
MI0000453	hsa-mir-135a-2	✓			
MI0000715	mmu-mir-135a-2	✓			
MI0000908	rno-mir-135a	✓			
MI0001169	gga-mir-135a-2	✓			
MI0001231	gga-mir-135a-1	✓			
MI0001264	gga-mir-135a-3	✓			
MI0002540	mml-mir-135a-2	✓			
MI0002541	ptr-mir-135a-2	✓			
MI0002542	ggo-mir-135a	✓			
MI0002543	ppy-mir-135a-2	✓			
MI0003388	fru-mir-135a	✓			
MI0003389	tni-mir-135a	✓			
MI0003692	dre-mir-135a	✓			
MI0005297	mdo-mir-135a	✓			
MI0006902	oan-mir-135a	✓			
MI0007624	mml-mir-135a-1	✓			
MI0008025	cfa-mir-135a-2	✓			
MI0008536	ptr-mir-135a-1	✓			
MI0009736	bta-mir-135a-2	✓			
MI0009737	bta-mir-135a-1	✓			

Search Results

Found **40** Homo sapiens miRNAs on chromosome chr13

Accession	ID	Chromosome	Start	End	Strand
MI0016862	hsa-mir-4499	chr13	20433778	20433846	-
MI0011282	hsa-mir-2276	chr13	24162416	24162504	+
MI0015835	hsa-mir-4305	chr13	39664034	39664135	-
MI0008190	hsa-mir-320d-1	chr13	40727828	40727875	-
MI0003635	hsa-mir-621	chr13	40810766	40810861	+
MI0014199	hsa-mir-3168	chr13	41101019	41101100	-
MI0017873	hsa-mir-5006	chr13	41568286	41568395	-
MI0025915	hsa-mir-8079	chr13	44196129	44196200	-
MI0016003	hsa-mir-3613	chr13	49996415	49996501	-
MI0000070	hsa-mir-16-1	chr13	50048973	50049061	-
MI0000069	hsa-mir-15a	chr13	50049119	50049201	-
MI0019300	hsa-mir-5693	chr13	51348567	51348639	-
MI0017336	hsa-mir-4703	chr13	51552589	51552667	+
MI0004065	hsa-mir-759	chr13	52810050	52810140	+
MI0006358	hsa-mir-1297	chr13	54311972	54312048	-
MI0017874	hsa-mir-5007	chr13	55174454	55174548	+
MI0014200	hsa-mir-3169	chr13	61199798	61199880	-
MI0016833	hsa-mir-548x-2	chr13	65966330	65966429	-
MI0017337	hsa-mir-4704	chr13	66218250	66218324	+



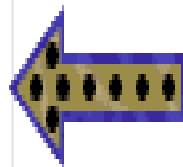


- Gene-miRNA Targets
- MicroRNA-gene Targets**
- MicroRNA-ncRNA Targets
- Gene-miRNA-pathway Targets
- Gene-miRNA-GO Targets
- Gene Class Targets
- Chromosome Targets
- Gene-miRNA-OMIM Targets
- Disease search
- Human Phenotype Ontologies
- Mitochondrial Targets
- Genomic location search
- Customized data set
- Holistic view

NEWS

- Gene-miRNA Targets
- MicroRNA-gene Targets
- Gene-miRNA-pathway Targets
- Gene-miRNA-GO Targets
- Gene Class Targets
- Cell line-based miRNAs
- Organ-based miRNAs
- miRNA-based search in organs
- Gene-miRNA-OMIM Targets
- Disease-ontology search
- Human Phenotype Ontologies
- Disease-based miRNAs**
- Literature on miRNAs
- miRNA processing proteins

NEWS





miRNA

Target mRNA

miRNA Expression

Downloads

FAQ

microRNA.org Releases:

- August 2010 release:
<http://www.microma.org/>
- September 2008 release:
(for a limited time):
<http://cbio.mskcc.org/microma-previous/>
- 2005 release:
<http://cbio.mskcc.org/mimaviewer/>

You are currently searching:

- All Species

miRNA Stats:

- Homo sapiens: 1100
- Mus musculus: 717
- Rattus norvegicus: 387
- Drosophila melanogaster: 186
- Caenorhabditis elegans: 233

Other microRNA resources:

- [PicTar](#)
- [TargetScan](#)
- [MicroCosm](#)
- [miRBase](#)

miRNA Search

miRNA(s)

miR-135a

☐ fuzzy search ("let-7" will match "let-7a", "let-7b", etc.)

Species:

Homo sapiens

Go



You are currently searching:

• Homo sapiens
• miR-135a

miRNA Stats:

• Homo sapiens: 1100
• Mus musculus: 717
• Rattus norvegicus: 387
• Drosophila melanogaster: 186
• Caenorhabditis elegans: 233

Searched for: miR-135a AND Homo sapiens

1 match [select all](#) [clear all](#)

miRNA	Genes Targeted	Links
<input type="checkbox"/> hsa-miR-135a	7,414	[view targets] [view expression profile] [view in miRBase] [view in miRò]

[[view mRNAs targeted by ALL selected miRNAs](#)] [[view expression profiles of ALL selected miRNAs](#)]

You are currently searching:

Homo sapiens
hsa-miR-135a
[[view targets](#)]
[[view expression profile](#)]

miRNA Stats:

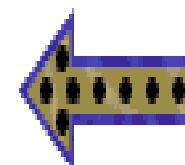
Homo sapiens: 1100
Mus musculus: 717
Rattus norvegicus: 387
Drosophila melanogaster: 186
Caenorhabditis elegans: 233

Selected miRNAs: hsa-miR-135a

Click [+](#) to view alternative isoforms.

Displaying results 1 - 50 of 7,414

[first](#) | [next](#) | [last](#)



mRNAs Targeted by hsa-miR-135a

mirSVR score

C9orf82	hsa	NM_024828,AK300739	[alignment details]	-2.97
+ B3GALT1	hsa	NM_194318,BX647932	[alignment details]	-2.60
- MYNN	hsa	NM_018657,AB079778,AF148848,AF155508,AK304281	[alignment details]	
For this gene, only alternative isoforms met search criteria				
		AY514901	[alignment details]	-2.59
TSPAN31	hsa	NM_005981,AK294128,BX647402	[alignment details]	-2.56
MAN2A1	hsa	NM_002372	[alignment details]	-2.33
- SUN1	hsa	NM_025154,NM_001130965,AK025737,AK302896,AK304150,BX648893...	[alignment details]	
For this gene, only alternative isoforms met search criteria				
		AK022469	[alignment details]	-2.27
- MFN1	hsa	NM_033540,AB043588,U95822	[alignment details]	
For this gene, only alternative isoforms met search criteria				
		AK000700	[alignment details]	-2.18
+ GULP1	hsa	NM_016315,AK298626,AK314498,BC068525	[alignment details]	-2.17
+ LCOR	hsa	NM_032440,AB058698,AL832106	[alignment details]	-2.11
+ FAM72D	hsa	NM_207418	[alignment details]	-2.09

miRNA.org

Targeting microRNAs to specific tissues



IMPORTANT NOTE:

This site is no longer supported. The **new and improved Web interface** can be accessed through [here](#).

WELCOME TO THE DIANA LAB WEBPAGE

Lab Intro

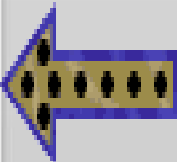
Computational predictive models are a key element of current systems biology. The focus of the DIANA lab is on the development of algorithms, databases and tools for interpreting and archiving genomic data in the framework of a systemic analysis. Current emphasis is on the analysis of **microRNA** (miRNA) and protein coding genes. MiRNAs are recently identified to be very abundant in mammalian organisms and play a key role in regulating development.

Comprehensive models work and integrate data at various levels of biological detail. Therefore the activities of the DIANA lab range from the analysis of expression regulation from deep sequencing data, the annotation of miRNA regulatory elements and targets to the interpretation of the role of miRNAs in various diseases.

Currently active projects of the DIANA-lab group include:

MicroRNA target prediction

DIANA-microT-CDS (v5.0): DIANA-microT-CDS is the 5th version of the microT algorithm. It is specifically trained on a positive and a negative set of miRNA Recognition Elements (MREs) located in both the 3'-UTR and CDS regions. DIANA-microT-CDS provides a significant increase in sensitivity compared to the previous version (65% vs 52%), when compared against experimental proteomics data. It exhibited the highest sensitivity at any level of specificity, when compared against other state of the art implementations. DIANA-microT-CDS has been updated (July 2012) to miRBase 18 and is fully compatible with the new miRNA nomenclature introduced in this version. It also provides hyperlinks to on-line servers such as iHOP and expression data for the selected



TarBase v.5c [Latest version of TarBase HERE](#)

[Download TarBase](#)

MORE RECENT VERSION OF TARBASE AVAILABLE!

The new and improved web interface containing updated data can be accessed through [here](#).

If used, please cite: The database of experimentally supported targets: a functional update of TarBase. (Papadopoulos GL, Reczko M, Simossis VA, Sethupathy P, Hatzigeorgiou AG.), Nucleic Acids Res. 2009 Jan;37(Database issue):D155-8. Epub 2008 Oct 27.

For issues regarding this application contact [maragkakis\[at\]fleming.gr](mailto:maragkakis[at]fleming.gr)

Select properties in the fields if you want to restrict your query | [Toggle More Options \(on/off\)](#) | [Start a new search](#)

Human

Any miRNA

HIC



[Target Search](#)
[Target Mining](#)
[Custom Prediction](#)
[FuncMir Collection](#)
[Data Download](#)
[Statistics](#)
[Help | FAQ](#)
[Comments](#)
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Choose one of the following search options:

Search by miRNA name

Human ▼

Go

Clear

Search by gene target

Human ▼

Gene Symbol ▼

Go

Clear

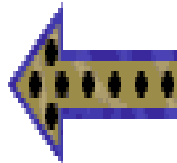
miRDB is an online database for miRNA target prediction and functional annotations. All the targets were predicted by a bioinformatics tool, MirTarget, which was developed by analyzing thousands of miRNA-target interactions from high-throughput sequencing experiments. Common features associated with miRNA target binding have been identified and used to predict miRNA targets with machine learning methods. miRDB hosts predicted miRNA targets in five species: human, mouse, rat, dog and chicken. As a recent update, users may provide their own sequences for customized target prediction.

Reference:

Nathan Wong and Xiaowei Wang (2015) miRDB: an online resource for microRNA target prediction and functional annotations. [Nucleic Acids Research](#), 43(D1):D146-152.

There are 158 predicted targets for hsa-miR-135a-3p in miRDB.

Target Detail	Target Rank	Target Score	miRNA Name	Gene Symbol	Gene Description
Details	1	98	hsa-miR-135a-3p	GLYCTK	glycerate kinase
Details	2	96	hsa-miR-135a-3p	PPP2R2B	protein phosphatase 2, regulatory subunit B, beta
Details	3	95	hsa-miR-135a-3p	CHD6	chromodomain helicase DNA binding protein 6
Details	4	92	hsa-miR-135a-3p	EP300	E1A binding protein p300
Details	5	90	hsa-miR-135a-3p	RHOBTB3	Rho-related BTB domain containing 3
Details	6	89	hsa-miR-135a-3p	PAIP1	poly(A) binding protein interacting protein 1
Details	7	89	hsa-miR-135a-3p	QSER1	glutamine and serine rich 1
Details	8	88	hsa-miR-135a-3p	COA6	cytochrome c oxidase assembly factor 6 homolog (S. cerevisiae)
Details	9	88	hsa-miR-135a-3p	FANCF	Fanconi anemia, complementation group F
Details	10	88	hsa-miR-135a-3p	VEZF1	vascular endothelial zinc finger 1
Details	11	87	hsa-miR-135a-3p	XG	Xg blood group
Details	12	85	hsa-miR-135a-3p	HGF	hepatocyte growth factor (hepapoietin A; scatter factor)
Details	13	84	hsa-miR-135a-3p	PTBP3	polypyrimidine tract binding protein 3
Details	14	84	hsa-miR-135a-3p	LCP1	lymphocyte cytosolic protein 1 (L-plastin)
Details	15	84	hsa-miR-135a-3p	ZBTB41	zinc finger and BTB domain containing 41
Details	16	83	hsa-miR-135a-3p	TFE3	transcription factor binding to IGHM enhancer 3
Details	17	83	hsa-miR-135a-3p	HIRA	histone cell cycle regulator
Details	18	82	hsa-miR-135a-3p	ZKSCAN1	zinc finger with KRAB and SCAN domains 1
Details	19	82	hsa-miR-135a-3p	C15orf53	chromosome 15 open reading frame 53
Details	20	82	hsa-miR-135a-3p	GAB1	GRB2-associated binding protein 1
Details	21	81	hsa-miR-135a-3p	ARL14	ADP-ribosylation factor-like 14
Details	22	79	hsa-miR-135a-3p	RERG	RAS-like, estrogen-regulated, growth inhibitor
Details	23	79	hsa-miR-135a-3p	SEL1L	sel-1 suppressor of lin-12-like (C. elegans)
Details	24	78	hsa-miR-135a-3p	ZSWIM6	zinc finger, SWIM-type containing 6
Details	25	77	hsa-miR-135a-3p	SSX2IP	synovial sarcoma, X breakpoint 2 interacting protein



Gene Expression Omnibus



GEO is a public functional genomics data repository supporting MIAME-compliant data submissions. Array- and sequence-based data are accepted. Tools are provided to help users query and download experiments and curated gene expression profiles.

Getting Started

[Overview](#)

[FAQ](#)

[About GEO DataSets](#)

[About GEO Profiles](#)

[About GEO2R Analysis](#)

[How to Construct a Query](#)

[How to Download Data](#)

Tools

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[Search for Gene Expression at GEO Profiles](#)

[Search GEO Documentation](#)

[Analyze a Study with GEO2R](#)

[GEO BLAST](#)

[Programmatic Access](#)

[FTP Site](#)

There are 27 results for "mir-135a" in the GEO DataSets Database.
There are 2 results for "mir-135a" in the GEO Profiles Database.

DataSets: 3848

Series: 62415

Platforms: 15114

Samples: 1613398

Information for Submitters

[Login to Submit](#)

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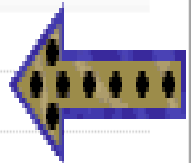
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[Citing and Linking to GEO](#)

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[GEO Publications](#)



miRNAMap

[Home](#)[About](#)[Browse](#)[Search](#)[Statistics](#)[Downloads](#)[Tutorial](#)keyword in ☒ microRNA or ☐ target gene :

Welcome to miRNAMap!

What's new?



More species



More prediction tools



Gene group search



miRNA target accessibility



faster and More comprehensive

Release 2.0 : July 2007



[Previous version - miRNAMap 1.0](#)

mature miRNA count:

Homo sapiens	542	<div></div>
Mus musculus	424	<div></div>
Danio rerio	371	<div></div>
Rattus norvegicus	261	<div></div>
Xenopus tropicalis	196	<div></div>
Gallus gallus	162	<div></div>
Caenorhabditis elegans	135	<div></div>
Fugu rubripes	133	<div></div>
Monodelphis domestica	111	<div></div>
Drosophila melanogaster	85	<div></div>
Anopheles gambiae	38	<div></div>
Canis familiaris	6	<div></div>

Citing miRNAMap2:

Hsu SD, Chu CH, Tsou AP, Chen SJ, Chen HC, Hsu PW, Wong YH, Chen YH, Chen GH, Huang HD (2008) 'miRNAMap 2.0: genomic maps of microRNAs in metazoan genomes.' Nucleic Acids Res. 2008 Jan;36(Database issue):D165-9. [\[Pubmed\]](#)

keyword in ☒ microRNA or ☐ target gene :

Browse

Choose one
species



Rattus norvegicus



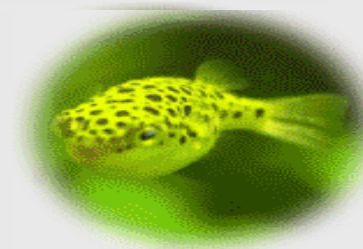
Homo sapiens



Mus musculus



Danio rerio



Fugu rubripes



Canis familiaris



Anopheles gambiae



Drosophila melanogaster



Caenorhabditis elegans



Monodelphis domestica



Xenopus tropicalis



Gallus gallus

The advancements and new features miRNAMap 2.0.

Features	miRNAMap 1.0	miRNAMap 2.0
Known miRNAs	miRBase (version 6.0)	miRBase (version 9.2)
Supported species	human, mouse, rat and dog	2 insects, 9 vertebrates and 1 worm
Experimental miRNA targets	Surveying literature	TarBase and Surveying literature
miRNA expression profiling	Lu. et al miRNA profiling in human	Lu. et al miRNA profiling in human Q-PCR miRNA profiling in human
Expression profiles of miRNA targets	-	NCBI-GEO-GDS596 (76 human tissues)
miRNA target prediction tools	miRanda	miRanda, RNAhybrid and TargetScan
Criteria for filtering the predicted miRNA targets	-	Criterion 1: <i>predicted by at least two tools</i> Criterion 2: <i>target genes contained multiple sites</i> Criterion 3: <i>target site is accessible</i>
Accessible region of miRNA target sites	-	Sfold
Tissue specificity of human miRNAs	-	Q-PCR miRNA profiling (18 human tissues)

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keyword in ☐ microRNA or ☒ target gene :

Items 1-2 of 2 displayed.

Show: Page 1 of 1

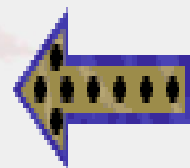
[Download Results \(Excel format\)](#)

ID	Species	Locate on chromosome	Locate on gene exon/intron/UTR	Get seq
hsa-mir-135a-1	Homo sapiens	3 : 52303275-52303364 : -	<input checked="" type="radio"/> OTTHUMT00000264972 (intron)	<input type="checkbox"/>
hsa-mir-135a-2	Homo sapiens	12 : 96481721-96481820 : +	<input checked="" type="radio"/> OTTHUMT00000262257 (intron)	<input type="checkbox"/>
				Select ALL <input type="checkbox"/>

Get miRNA's sequence:



تقدیر برای مرض تشکر





References

1. Singh SR, Rameshwar P. **MicroRNA in Development and in the Progression of Cancer**: Springer; 2014.
2. Selbach M, Schwanhäusser B, Thierfelder N, Fang Z, Khanin R, Rajewsky N. **Widespread changes in protein synthesis induced by microRNAs**. Nature. 2008;455(7209):58-63.
3. Ferguson EL, Sternberg PW, Horvitz HR. **A genetic pathway for the specification of the vulval cell lineages of Caenorhabditis elegans**. Nature. 1987;326(6110):259-67.
4. Ruvkun G, Giusto J. **The Caenorhabditis elegans heterochronic gene lin-14 encodes a nuclear protein that forms a temporal developmental switch**. Nature. 1989;338(6213):313-9.



5. Lee RC, Feinbaum RL, Ambros V. **The C. elegans heterochronic gene lin-4 encodes small RNAs with antisense complementarity to lin-14.** cell. 1993;75(5):843-54.
6. Parvini N, Ahmadi S. **Role of MicroRNAs in Development of Immune Cells and Nervous System and their Relation to Multiple Sclerosis.** The Neuroscience Journal of Shefaye Khatam. 2015;3(1):0.
7. Calin GA, Dumitru CD, Shimizu M, Bichi R, Zupo S, Noch E, et al. **Frequent deletions and down-regulation of micro-RNA genes miR15 and miR16 at 13q14 in chronic lymphocytic leukemia.** Proceedings of the National Academy of Sciences. 2002;99(24):15524-9.
8. Garzon R, Calin GA, Croce CM. **MicroRNAs in cancer.** Annual review of medicine. 2009;60:167-79.



9. Saito Y, Liang G, Egger G, Friedman JM, Chuang JC, Coetzee GA, et al. **Specific activation of microRNA-127 with downregulation of the proto-oncogene BCL6 by chromatin-modifying drugs in human cancer cells.** Cancer cell. 2006;9(6):435-43
10. Ferracin M, Lupini L, Salamon I, Saccenti E, Zanzi MV, Rocchi A, et al. **Absolute quantification of cell-free microRNAs in cancer patients.** Oncotarget. 2015;6(16):14545-55.
11. Zhao Y, Samal E, Srivastava D. **Serum response factor regulates a muscle-specific microRNA that targets Hand2 during cardiogenesis.** Nature. 2005;436(7048):214-20
12. van Rooij E, Sutherland LB, Liu N, Williams AH, McAnally J, Gerard RD, Richardson JA, Olson EN. **A signature pattern of stress-responsive microRNAs that can evoke cardiac hypertrophy and heart failure.** Proc Natl Acad Sci USA. 2006;103(48):18255-60.



13. Edwards JK, Pasqualini R, Arap W, Calin GA. **MicroRNAs and ultraconserved genes as diagnostic markers and therapeutic targets in cancer and cardiovascular diseases.** J Cardiovasc Transl Res. 2010;3(3):271-9
14. Bian S, Sun T. **Functions of noncoding RNAs in neural development and neurological diseases.** Molecular neurobiology. 2011;44(3):359-73
15. Zhou C, Gu H, Fan R, Wang B, Lou J. **MicroRNA 302/367 Cluster Effectively Facilitates Direct Reprogramming from Human Fibroblasts into Functional Neurons.** Stem Cells Dev. 2015 ;2015.
16. Möbus S, Yang D, Yuan Q, Lüdtke TH-W, Balakrishnan A, Sgodda M, et al. **MicroRNA-199a-5p inhibition enhances the liver repopulation ability of human embryonic stem cell-derived hepatic cells.** Journal of hepatology. 2015;62(1):101-10.
17. Chim SS, Shing TK, Hung EC, Leung TY, Lau TK, Chiu RW, Lo YM. **Detection and characterization of placental microRNAs in maternal plasma.** Clin Chem. 2008;54(3):482-90



18. Lawrie CH, Gal S, Dunlop HM, Pushkaran B, Liggins AP, Pulford K, Banham AH, Pezzella F, Boulwood J, Wainscoat JS, Hatton CS, Harris AL. **Detection of elevated levels of tumour-associated microRNAs in serum of patients with diffuse large B-cell lymphoma.** Br J Haematol. 2008;141(5):672-5
19. Murata K, Yoshitomi H, Tanida S, Ishikawa M, Nishitani K, Ito H, Nakamura T. **Plasma and synovial fluid microRNAs as potential biomarkers of rheumatoid arthritis and osteoarthritis.** Arthritis Res Ther. 2010;12(3):R86
20. Tsochandaridis M, Nasca L, Toga C, Levy-Mozziconacci A. **Circulating microRNAs as clinical biomarkers in the predictions of pregnancy complications.** BioMed research international. 2015;2015.
21. Tian T, Wang J, Zhou X. **A review: microRNA detection methods.** Organic & biomolecular chemistry. 2015;13(8):2226-38.